

## **EXHIBIT A**

```
#!/internet/bin/perl5.002 -w
# Copyright (c) 1998
        Eugene Wang
# *** BEGIN ***
#-----
#input sequence (File 0) to compare
#-----
if ($#ARGV < 2) {die "argv < 2";}
open(EnzymeInput, $ARGV[0]) || die "Cannot open input file $ARGV[0]";
#print "Input Enzyme 1 sequence = ";
$Elsequence = <EnzymeInput>;
chomp $Elsequence;
$lenE1Seq = length($E1sequence);
$Elsequence =~ tr/a-z/A-Z/;
$E1ExtLoc = <EnzymeInput>;
chomp($E1ExtLoc);
$lenElTotal = $lenElSeq + $ElExtLoc;
#print "Input Enzyme 2 sequence = ";
$E2sequence = <EnzymeInput>;
chomp $E2sequence;
$E2sequence = reverse($E2sequence);
$lenE2Seq = length($E2sequence);
$E2sequence =~ tr/a-z/A-Z/;
$E2ExtLoc = <EnzymeInput>;
chomp($E2ExtLoc);
$lenE2Total = $lenE2Seq + $E2ExtLoc;
$lenE1Extra = $E2ExtLoc - $E1ExtLoc;
$E1SizeStart = <EnzymeInput>;
chomp($E1SizeStart);
$E1SizeEnd = <EnzymeInput>;
chomp($E1SizeEnd);
```

#open input FASTA file (File 1)

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#print "Input file name = ";
#$fname = <>;
#chomp $fname;
#$fname = "H DJ0167F23.seq";
open(Infile, $ARGV[1]) || die "Cannot open input file $ARGV[1]";
#open output file (File 2)
open (Outfile, ">$ARGV[2]") || die "Cannot open output file $ARGV[2]";
#open (Outfile,">output.txt");
#print Outfile "Qualifier\tSequence";
#read input FASTA file
                       #header line
$line = <Infile>;
print Outfile "$line";
$linecount = 0;
$FullSeq = "";
#check headerline format
chomp $line;
@fields = split (/\|/,$line);
ntokens = 0;
foreach (@fields) {$ntokens++;}
#$ntokens = @fields;
if ($ntokens > 3)
   {$FragmentID = $fields[3];}
else
   $line =~ s/^> />/;
   @fields = split (/ /,$line);
   sntokens = 0;
   foreach (@fields) {$ntokens++;}
    if ($ntokens > 0)
       {$FragmentID = $fields[0]; $FragmentID =~ s/^>//;}
    else
       {$FragmentID = "UnknownFragment";}
                                 #read in a line
 while ($line = <Infile>)
         {
```

```
print Outfile "Enzyme top strand: ";
print Outfile "(5\'-$Elsequence";
                  {print Outfile "(N)$E1ExtLoc";}
if ($E1ExtLoc>0)
print Outfile "-3\')";
print Outfile "\n";
print Outfile "Enzyme bottom strand: ";
print Outfile "(5\'-";
                    {print Outfile "(N)$E2ExtLoc";}
if ($E2ExtLoc>0)
print Outfile "$E2sequence-3\')";
print Outfile " or ";
my $ts = reverse($E2sequence);
print Outfile "(3\'-$ts";
                   {print Outfile "(N)$E2ExtLoc";}
if ($E2ExtLoc>0)
print Outfile "-5\')";
print Outfile "\n";
print Outfile "Segment size: $E1SizeStart - $E1SizeEnd\n";
$minLen = $lenE1Total < $lenE2Total ? $lenE1Total : $lenE2Total;</pre>
$maxLen = $lenE1Total > $lenE2Total ? $lenE1Total : $lenE2Total;
nMatchE1 = 0;
nSelected = 0;
@EnzLocLeft = ();
@EnzLocRight = ();
@EnzTypeLeft = ();
@EnzTypeRight = ();
if ($minLen > 0)
         for ($i=0; $i <= $lenFullSeq-$lenE1Seq; $i++)</pre>
        for ($i=0; $i <= $lenFullSeq-$maxLen; $i++)</pre>
            if (substr($FullSeq,$i,$lenE1Seq) eq $E1sequence)
                         $EnzLocLeft[$nMatchE1] = $i + $lenE1Total;
##have to use push()
                         $EnzTypeLeft[$nMatchE1] = 1;
                         push(@EnzLocLeft,$i + $lenElTotal);
                         push(@EnzTypeLeft,1);
                  print Outfile "$nMatchE1\t$i\t";
                  print Outfile "type 1\t";
                         print Outfile "$Elsequence\t";
                         print Outfile substr($FullSeq,$i,$lenElTotal);
                  print Outfile "\n";
                         if (nMatchE1 > 0)
                            push(@EnzLocRight,$i + $lenElTotal-1);
                            push(@EnzTypeRight,1);
```

```
$nMatchE1++;
                if (substr($FullSeq,$i+$E2ExtLoc,$lenE2Seq) eq
$E2sequence)
                elsif (substr($FullSeq,$i+$E2ExtLoc,$lenE2Seq) eq
$E2sequence)
                  {
                         $EnzLocLeft[$nMatchE1] = $i;
                         $EnzCutLeft[$nMatchE1] = 2;
                         push(@EnzLocLeft,$i);
                         push(@EnzTypeLeft,2);
                  print Outfile "$nMatchE1\t$i\t";
                  print Outfile "type 2\t";
                  print Outfile "$E2sequence\t";
                         print Outfile substr($FullSeq,$i,$lenE2Total);
                  print Outfile "\n";
                         if (\$nMatchE1 > 0)
                            push(@EnzLocRight,$i-1);
                            push(@EnzTypeRight,2);
                         $nMatchE1++;
                   }
             }
        if ($nMatchE1 > 0)
           push(@EnzLocRight,$i-1);
           push(@EnzTypeRight,2);
        print Outfile "Number of segments: $nMatchEl\n";
        if ($nMatchE1 != ($#EnzLocRight+1)) {die ("Counting
error...nMatchE1($nMatchE1) != $#EnzLocRight");}
        print Outfile "Matched loci:\n";
         for ($i=0; $i < $nMatchE1; $i++)</pre>
                 print Outfile "$EnzLocLeft[$i]\t";
         print Outfile "\nSegment Size:\n";
         for ($i=0; $i < $nMatchE1-1; $i++)</pre>
                 $tmpSegSize = $EnzLocRight[$i] - $EnzLocLeft[$i] + 1;
                 if ($tmpSegSize >= $E1SizeStart && $tmpSegSize <=</pre>
$E1SizeEnd)
                          $SeqSelected[$nSelected++] = $i;
```

```
print Outfile "$tmpSegSize\t";
      print out the Segment (E1) sequences
print Outfile "\nSegments Selected ($nSelected):";
for ($i=0; $i < $nSelected; $i++)
        $selSeq = $SegSelected($i);
        $E1left = $EnzLocLeft[$selSeq];
        $E1right = $EnzLocRight[$selSeq];
        if ($lenE1Extra > 0) {$E1right += $lenE1Extra;}
        else {$Elleft += $lenElExtra;}
        $lenSelSeq = $Elright - $Elleft + 1;
        $OutputHeaderLine = ">" . $FragmentID ."_" .$selSeq . "\tsize=" .
$lenSelSeq;
        $OutputHeaderLine .= "\tLoci=" . $Elleft . "-" . $Elright;
        $OutputHeaderLine .= "\tEnz$EnzTypeLeft[$selSeq]-
Enz$EnzTypeRight[$selSeq]";
        print Outfile "\n$OutputHeaderLine";
        print "$OutputHeaderLine";
        Segment sequence
        $SeqEltoNextE1 = substr($FullSeq,$Elleft,$lenSelSeq);
        print Outfile "\n$SeqEltoNextEl\n";
        print "\n$SeqEltoNextEl\n";
return ($lenFullSeq);
```





## **EXHIBIT B**

```
#!/internet/bin/perl5.002 -w
#*********
# Copyright (c) 1998
# Author: Eugene Wang
# Title: Ligate
# Purpose: Find matching segments/sequences in two files
#****
if ($#ARGV != 2) {die "Number of argv ($#ARGV+1) != 3";}
#input file
open(InfileLigate, $ARGV[0]) or die "Open error...$ARGV[0]\n";
$locLigate = <InfileLigate>;
chomp $locLigate;
$seqLigate = <InfileLigate>;
chomp $seqLigate;
close (InfileLigate);
#-----
#-----
open(Infile, $ARGV[1]) or die "Open error...$ARGV[1] \n";
$OutName = $ARGV[2];
open (Outfile, ">$OutName") or die("Open error...$OutName");
$alreadyReadOne = 0;
$sequence = "";
while ($line = <Infile>)  #read in a line
      chomp $line;
       next if ($line eq "");
                                       ##if first char is a '#'
       if ($line =~ /^#/ || $line =~ /^>/)
or '>'
         if ($alreadyReadOne == 1) {
             if (&Ligate($sequence,$locLigate,$seqLigate) == 1) {
                print Outfile "$headerLine\n";
                print Outfile "$sequence\n";
                };
             $sequence = "";
```





```
$headerLine = $line;
         $alreadyReadOne = 1;
        . }
      else
         $sequence .= $line;
      }
if ($alreadyReadOne == 1) {
      if (&Ligate($sequence,$locLigate,$seqLigate) == 1) {
         print Outfile "$headerLine\n";
         print Outfile "$sequence\n";
         };
      }
close (Infile);
close (Outfile);
#compare sequence with Ligation Adapter sequence
############
sub Ligate()
local $retcode = 0;
local ($seq,$locLigate,$seqLigate) = @_;
local $lenLigate = length($seqLigate);
local $lenSeq = length($seq);
if ((substr($seq,$locLigate,$lenLigate) eq $seqLigate) &&
   (substr($seq,$lenSeq-$locLigate-$lenLigate,$lenLigate) eq
$seqLigate)) {
  $retcode = 1;
  }
return $retcode;
```